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(FILE 'HOME' ENTERED AT 12:13:28 ON 04 MAR 2004)

FILE 'REGISTRY' ENTERED AT 12:13:41 ON 04 MAR 2004

L1 1 S 193270-76-7

L2 1 S 193272-70-7

FILE 'CAPLUS' ENTERED AT 12:15:27 ON 04 MAR 2004

L3 13 S L1 OR L2

L4 1 S L3 AND (APPETITE OR FOOD OR HUNGER OR DEPRESS?)

FILE 'MEDLINE' ENTERED AT 12:39:37 ON 04 MAR 2004

L5 52147 S GROWTH(3A)HORMONE

L6 772 S L5(L) (APPETITE OR HUNGER OR FOOD)

L7 31 S L5(S)APPETITE

L8 17 S L7 NOT PY>=2000

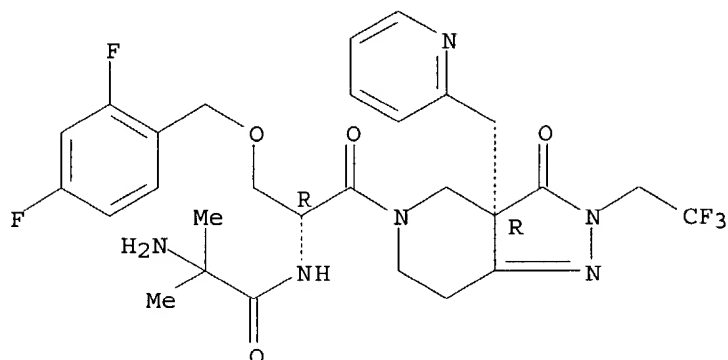
L9 38 S L5(L) (APPETITE(10A) (INCREAS? OR STIMULAT? ))

L10 15 S L9 NOT PY>=2000

*Instant Compound*

L2 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 193272-70-7 REGISTRY

Absolute stereochemistry.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

CN Propanamide, 2-amino-N-[(1R)-1-[[2,4-difluorophenyl)methoxy)methyl]-2-[(3aR)-2,3,3a,4,6,7-hexahydro-3-oxo-3a-(2-pyridinylmethyl)-2-(2,2,2-trifluoroethyl)-5H-pyrazolo[4,3-c]pyridin-5-yl]-2-oxoethyl]-2-methyl-(9CI) (CA INDEX NAME)

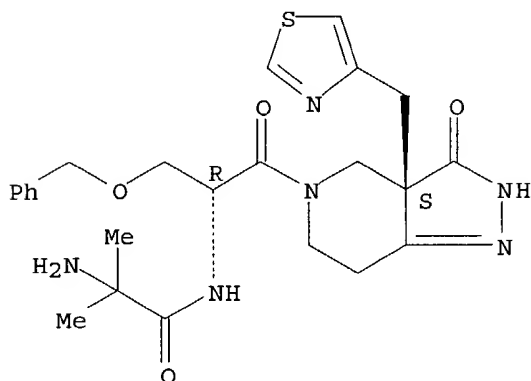
OTHER CA INDEX NAMES:

CN Propanamide, 2-amino-N-[1-[[2,4-difluorophenyl)methoxy)methyl]-2-[2,3,3a,4,6,7-hexahydro-3-oxo-3a-(2-pyridinylmethyl)-2-(2,2,2-trifluoroethyl)-5H-pyrazolo[4,3-c]pyridin-5-yl]-2-oxoethyl]-2-methyl-, [R-(R\*,R\*)]-

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 193270-76-7 REGISTRY

Absolute stereochemistry.

*Instant compound*



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

CN Propanamide, 2-amino-N-[(1R)-2-[(3aS)-2,3,3a,4,6,7-hexahydro-3-oxo-3a-(4-thiazolylmethyl)-5H-pyrazolo[4,3-c]pyridin-5-yl]-2-oxo-1-[(phenylmethoxy)methyl]ethyl]-2-methyl- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Propanamide, 2-amino-N-[2-[2,3,3a,4,6,7-hexahydro-3-oxo-3a-(4-thiazolylmethyl)-5H-pyrazolo[4,3-c]pyridin-5-yl]-2-oxo-1-[(phenylmethoxy)methyl]ethyl]-2-methyl-, [R-(R\*,S\*)]-

L8 ANSWER 10 OF 17 MEDLINE on STN  
 ACCESSION NUMBER: 95274367 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 7754750  
 TITLE: Childhood obesity: pathophysiology and treatment.  
 AUTHOR: Klish W J  
 CORPORATE SOURCE: Department of Pediatric Nutrition and Gastroenterology,  
 Baylor College of Medicine, Texas Children's Hospital,  
 Houston 77030, USA.  
 SOURCE: Acta paediatrica Japonica; Overseas edition, (1995 Feb) 37  
 (1) 1-6. Ref: 18  
 Journal code: 0370357. ISSN: 0374-5600.  
 PUB. COUNTRY: Australia  
 DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
 General Review; (REVIEW)  
 (REVIEW LITERATURE)  
 LANGUAGE: English  
 FILE SEGMENT: Priority Journals  
 ENTRY MONTH: 199506  
 ENTRY DATE: Entered STN: 19950629  
 Last Updated on STN: 19950629  
 Entered Medline: 19950619

AB Childhood obesity is among the most difficult problems which pediatricians treat. It is frequently ignored by the pediatrician or viewed as a form of social deviancy, and blame for treatment failure placed on the patients or their families. The definition of obesity is difficult. Using total body electrical conductivity (TOBEC) technology, total body fat ranges between 12% and 30% of total body weight in normal children and adolescents. This is influenced not only by age, but also by physical fitness. Anthropometry is the easiest way to define obesity. Children whose weight exceeds 120% of that expected for their height are considered overweight. Skinfold thickness and body mass index are indices of obesity that are more difficult to apply to the child. Childhood obesity is associated with obese parents, a higher socioeconomic status, increased parental education, small family size and a sedentary lifestyle. Genetics also clearly plays a role. Studies have demonstrated that obese and non-obese individuals have similar energy intakes implying that obesity results from very small imbalances of energy intake and expenditure. An excess intake of only 418 kJ per day can result in about 4.5 kg of excess weight gain per year. Small differences in basal metabolic rate or the thermic effects of food may also account for the difference in energy balance between the obese and non-obese. In the Prader Willi Syndrome, there appears to be a link between appetite and body fatness. When placed on growth hormone, lean body mass increases, body fat decreases, sometimes to normal, and appetite becomes more normal. (ABSTRACT TRUNCATED AT 250 WORDS)

09/893, 014

L10 ANSWER 3 OF 15 MEDLINE on STN  
ACCESSION NUMBER: 2000036825 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 10567856  
TITLE: Co-localization of growth hormone secretagogue receptor and  
NPY mRNA in the arcuate nucleus of the rat.  
AUTHOR: Willesen M G; Kristensen P; Romer J  
CORPORATE SOURCE: Department of Histology, Health Care Pharmacology, Health  
Care Discovery, Novo Nordisk A/S, Bagsvaerd, Denmark.  
SOURCE: Neuroendocrinology, (1999 Nov) 70 (5) 306-16.  
Journal code: 0035665. ISSN: 0028-3835.  
PUB. COUNTRY: Switzerland  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200004  
ENTRY DATE: Entered STN: 20000427  
Last Updated on STN: 20000427  
Entered Medline: 20000414

AB | **Growth hormone** secretagogues (GHS) are small,  
synthetic compounds which have the potential of releasing **growth  
hormone** (GH) from the pituitary. The mechanism of action of GHS  
has not been fully elucidated. A specific GHS receptor (GHS-R) is  
expressed in the pituitary gland and in several areas of the brain  
including the hypothalamus. We have characterized the  
GHS-R-mRNA-expressing neurons with respect to co-expression of selected  
neurotransmitters in the hypothalamus. This was done by dual chromogenic  
and autoradiographic in situ hybridization with riboprobes for GHS-R mRNA  
and neuropeptide Y (NPY), pro-opiomelanocortin (POMC), somatostatin (SRIH)  
or GH-releasing hormone (GHRH) mRNA. In the arcuate nucleus, GHS-R mRNA  
was expressed in 94 +/- 1% of the neurons expressing NPY, 8 +/- 2% of  
those expressing POMC and 30 +/- 6% expressing SRIH mRNA. 20-25% of the  
GHRH- mRNA-expressing neurons contained GHS-R mRNA, whereas the vast  
majority of the arcuate GHS-R-mRNA-containing cells did not contain GHRH  
mRNA. The finding of a significant co-expression of GHS-R and NPY mRNA in  
the arcuate nucleus is in accordance with the previous demonstration by  
Dickson et al. that c-Fos is induced in NPY neurons following GHS  
administration. These results indicate that GHS have other effects on  
neuroendocrine regulation than GH release via GHRH neurons. Stimulation  
of the arcuate NPY neurons via GHS-R may explain the **increased  
appetite** and the cortisol release seen after administration of  
some GHS compounds.

09/893,014

L10 ANSWER 12 OF 15 MEDLINE on STN  
ACCESSION NUMBER: 87085987 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 3794875  
TITLE: Enhancement of linear growth and weight gain by  
cyproheptadine in children with hypopituitarism receiving  
growth hormone therapy.  
AUTHOR: Kaplowitz P B; Jennings S  
CONTRACT NUMBER: M01-RR-00065 (NCRR)  
SOURCE: Journal of pediatrics, (1987 Jan) 110 (1) 140-3.  
Journal code: 0375410. ISSN: 0022-3476.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: (CLINICAL TRIAL)  
Journal; Article; (JOURNAL ARTICLE)  
(RANDOMIZED CONTROLLED TRIAL)  
LANGUAGE: English  
FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals  
ENTRY MONTH: 198702  
ENTRY DATE: Entered STN: 19900302  
Last Updated on STN: 19980206  
Entered Medline: 19870205

- AB Cyproheptadine (Cp), an antihistamine serotonin antagonist drug with appetite-stimulating activity, was given to children with growth hormone (GH) deficiency to test the hypothesis that increased weight gain would enhance the effect of GH on linear growth. Six patients with idiopathic GH deficiency received GH 0.08 U/kg three times per week plus Cp 0.25 to 0.4 mg/kg/day for 4-month periods, alternating with 4-month periods of GH plus placebo, on average for 16 months. Overall, height velocity (HV) increased from 9.1 +/- 2.4 with GH alone to 12.1 +/- 2.1 cm/yr with GH-Cp (P = 0.01) and weight velocity (WV) increased substantially from 1.3 +/- 1.3 to 7.8 +/- 3.6 kg/yr (P = 0.01). For 10 of 11 8-month treatment intervals completed, HV was greater during GH-Cp treatment than during GH alone, and there was a good correlation between HV and WV for each 4-month observation period (r = 0.64, P less than 0.002). These findings should be considered preliminary because of the small number of patients, but suggest that weight gain induced by cyproheptadine results in improved linear growth in patients given GH and that this drug may be useful in optimizing the response to GH therapy.
- AB Cyproheptadine (Cp), an antihistamine serotonin antagonist drug with appetite-stimulating activity, was given to children with growth hormone (GH) deficiency to test the hypothesis that increased weight gain would enhance the effect of GH on linear growth. Six. . .